RAW SEQUENCE LISTING DATE: 11/19/2001 PATENT APPLICATION: US/09/965,807 TIME: 14:24:06

Input Set : N:\Crf3\RULE60\09965807.txt
Output Set: N:\CRF3\11192001\1965807.raw

SEQUENCE LISTING

```
(1) GENERAL INFORMATION:
             (i) APPLICANT: Matalon, Reuben
      6
                            Kaul, Rajinder
      7
                            Gao, Guang Ping
      8
                            Balamurugan, Kuppareddi
      9
                            Michals-Matalon, Kimberlee
     10
            (ii) TITLE OF INVENTION: Aspartoacylase Gene, Protein, and
     12
                                      Methods of Screening for Mutations Associated with
     13
Canavan
     14
           (iii) NUMBER OF SEQUENCES: 27
     16
            (iv) CORRESPONDENCE ADDRESS:
     18
                  (A) ADDRESSEE: Millen, White, Zelano & Branigan, P.C.
     19
                  (B) STREET: 2200 Clarendon Boulevard, Suite 1400
     20
                  (C) CITY: Arlington
     21
                                                                ENTERED
                  (D) STATE: Virginia
     22
                  (E) COUNTRY: U.S.A.
     23
     24
                  (F) ZIP: 22201
             (V) COMPUTER READABLE FORM:
     26
                  (A) MEDIUM TYPE: Floppy disk
     27
                  (B) COMPUTER: IBM PC compatible
     28
                  (C) OPERATING SYSTEM: PC-DOS/MS-DOS
     29
                  (D) SOFTWARE: PatentIn Release #1.0, Version #1.25
     30
            (vi) CURRENT APPLICATION DATA:
     32
                  (A) APPLICATION NUMBER: US/09/965,807
C--> 33
                  (B) FILING DATE: 01-Oct-2001
C-->34
                  (C) CLASSIFICATION:
     40
           (vii) PRIOR APPLICATION DATA:
     37
                   (A) APPLICATION NUMBER: US 08/128,020
     38
                   (B) FILING DATE: 29-SEP-1993
     39
          (viii) ATTORNEY/AGENT INFORMATION:
     42
                   (A) NAME: Hamlet-King, Diana
     43
                   (B) REGISTRATION NUMBER: 33,302
     44
                   (C) REFERENCE/DOCKET NUMBER: Shutt 1
     45
            (ix) TELECOMMUNICATION INFORMATION:
     47
                   (A) TELEPHONE: 703-243-6333
     48
                   (B) TELEFAX: 703-243-6410
     49
                   (C) TELEX: 64191
     50
     53 (2) INFORMATION FOR SEQ ID NO: 1:
             (i) SEQUENCE CHARACTERISTICS:
     55
                   (A) LENGTH: 1435 base pairs
     56
                   (B) TYPE: nucleic acid
     57
                   (C) STRANDEDNESS: double
     58
                   (D) TOPOLOGY: linear
     59
     62
            (ix) FEATURE:
                   (A) NAME/KEY: CDS
     63
```

(B) LOCATION: 159..1097

64

PATENT APPLICATION: US/09/965,807

DATE: 11/19/2001 TIME: 14:24:06

| 70 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1: | | | | | | | | | | | |
|---|---------------------|---|--|--|--|--|--|--|--|--|--|
| 72 TTGTAACAGA AAATTAAAAT ATACTCCACT CAAGGGAATT CTGTACTTTG CCC | CTTTTGGT 60 | | | | | | | | | | |
| 74 AAAGTCTCAT TTACATTTCT AAACCTTTCT TAAGAAAATC GAATTTCCTT TGA | ATCTCTCT 120 | | | | | | | | | | |
| 76 TCTGAATTGC AGAAATCAGA TAAAAACTAC TTGGTGAA ATG ACT TCT TGT | CAC 173 | | | | | | | | | | |
| | Not The Con Cyc Hic | | | | | | | | | | |
| 78 | | | | | | | | | | | |
| 80 ATT GCT GAA GAA CAT ATA CAA AAG GTT GCT ATC TTT GGA GGA AG | CC CAT 221 | | | | | | | | | | |
| 81 Ile Ala Glu Glu His Ile Gln Lys Val Ala Ile Phe Gly Gly Th | hr His | | | | | | | | | | |
| 82 10 15 | 20 | | | | | | | | | | |
| 84 GGG AAT GAG CTA ACC GGA GTA TTT CTG GTT AAG CAT TGG CTA GA | AG AAT 269 | | | | | | | | | | |
| 85 Gly Asn Glu Leu Thr Gly Val Phe Leu Val Lys His Trp Leu Gl | lu Asn | | | | | | | | | | |
| 86 25 30 35 | | | | | | | | | | | |
| 88 GGC GCT GAG ATT CAG AGA ACA GGG CTG GAG GTA AAA CCA TTT AT | IT ACT 317 | | | | | | | | | | |
| 89 Gly Ala Glu Ile Gln Arg Thr Gly Leu Glu Val Lys Pro Phe I | le Thr | | | | | | | | | | |
| 90 40 45 50 | | | | | | | | | | | |
| 92 AAC CCC AGA GCA GTG AAG AAG TGT ACC AGA TAT ATT GAC TGT GA | AC CTG 365 | | | | | | | | | | |
| 93 Asn Pro Arg Ala Val Lys Lys Cys Thr Arg Tyr Ile Asp Cys As | sp Leu | | | | | | | | | | |
| 94 55 60 65 | | | | | | | | | | | |
| 96 AAT CGC ATT TTT GAC CTT GAA AAT CTT GGC AAA AAA ATG TCA GA | AA GAT 413 | | | | | | | | | | |
| 97 Asn Arg Ile Phe Asp Leu Glu Asn Leu Gly Lys Lys Met Ser G | lu Asp | | | | | | | | | | |
| 98 70 75 80 | 85 | | | | | | | | | | |
| 100 TTG CCA TAT GAA GTG AGA AGG GCT CAA GAA ATA AAT CAT TTA | TTT GGT 461 | 1 | | | | | | | | | |
| 101 Leu Pro Tyr Glu Val Arg Arg Ala Gln Glu Ile Asn His Leu | Phe Gly | | | | | | | | | | |
| 102 | 100 | _ | | | | | | | | | |
| 104 CCA AAA GAC AGT GAA GAT TCC TAT GAC ATT ATT TTT GAC CTT | CAC AAC 509 | 9 | | | | | | | | | |
| 105 Pro Lys Asp Ser Glu Asp Ser Tyr Asp Ile Ile Phe Asp Leu I | His Asn | | | | | | | | | | |
| 106 105 110 115 | | _ | | | | | | | | | |
| 108 ACC ACC TCT AAC ATG GGG TGC ACT CTT ATT CTT GAG GAT TCC | AGG AAT 55' | / | | | | | | | | | |
| 109 Thr Thr Ser Asn Met Gly Cys Thr Leu Ile Leu Glu Asp Ser | Arg Asn | | | | | | | | | | |
| 110 120 125 130 | | _ | | | | | | | | | |
| 112 AAC TTT TTA ATT CAG ATG TTT CAT TAC ATT AAG ACT TCT CTG | GCT CCA 60! | 5 | | | | | | | | | |
| 113 Asn Phe Leu Ile Gln Met Phe His Tyr Ile Lys Thr Ser Leu | Ala Pro | | | | | | | | | | |
| 114 . 135 140 145 | mam | 2 | | | | | | | | | |
| 116 CTA CCC TGC TAC GTT TAT CTG ATT GAG CAT CCT TCC CTC AAA | TAT GCG 65 | 3 | | | | | | | | | |
| 117 Leu Pro Cys Tyr Val Tyr Leu Ile Glu His Pro Ser Leu Lys | Tyr Ala | | | | | | | | | | |
| 118 150 155 160 | 165 GGT CCT 70: | 1 | | | | | | | | | |
| 120 ACC ACT CGT TCC ATA GCC AAG TAT CCT GTG GGT ATA GAA GTT | | _ | | | | | | | | | |
| 121 Thr Thr Arg Ser Ile Ala Lys Tyr Pro Val Gly Ile Glu Val | GLY Pro | | | | | | | | | | |
| 122 170 175 | | ۵ | | | | | | | | | |
| 124 CAG CCT CAA GGG GTT CTG AGA GCT GAT ATC TTG GAT CAA ATG | | פ | | | | | | | | | |
| 125 Gln Pro Gln Gly Val Leu Arg Ala Asp Ile Leu Asp Gln Met | Arg Lys | | | | | | | | | | |
| 126 185 190 195 | GGA AAA 79° | 7 | | | | | | | | | |
| 128 ATG ATT AAA CAT GCT CTT GAT TTT ATA CAT CAT TTC AAT GAA | | • | | | | | | | | | |
| 129 Met Ile Lys His Ala Leu Asp Phe Ile His His Phe Asn Glu | ата пар | | | | | | | | | | |
| 130 200 | AAA GTT 84. | 5 | | | | | | | | | |
| 134 GAA TTT CCT CCC TGC GCC ATT GAG GTC TAT AAA ATT ATA GAG | | _ | | | | | | | | | |
| 135 Glu Phe Pro Pro Cys Ala Ile Glu Val Tyr Lys Ile Ile Glu | nlo Arr | | | | | | | | | | |
| 130 213 | CAT CCT 89 | 3 | | | | | | | | | |
| 138 GAT TAC CCC CGG GAT GAA AAT GGA GAA ATT GCT GCT ATC ATC | CAI CCI 09 | J | | | | | | | | | |

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| | Asp Tyr Pro Arg Asp Glu Asn Gly Glu Ile Ala Ala Ile Ile His Pro | | | | | | | | | |
|--|---|------|--|--|--|--|--|--|--|--|
| | 230 235 240 245 | | | | | | | | | |
| | AAT CTG CAG GAT CAA GAC TGG AAA CCA CTG CAT CCT GGG GAT CCC ATG | 941 | | | | | | | | |
| | Asn Leu Gln Asp Gln Asp Trp Lys Pro Leu His Pro Gly Asp Pro Met | | | | | | | | | |
| 144 | | 000 | | | | | | | | |
| | TTT TTA ACT CTT GAT GGG AAG ACG ATC CCA CTG GGC GGA GAC TGT ACC | 989 | | | | | | | | |
| | Phe Leu Thr Leu Asp Gly Lys Thr Ile Pro Leu Gly Gly Asp Cys Thr 265 270 275 | | | | | | | | | |
| 148 | 265 270 275 GTG TAC CCC GTG TTT GTG AAT GAG GCC GCA TAT TAC GAA AAG AAA GAA | 1037 | | | | | | | | |
| | Val Tyr Pro Val Phe Val Asn Glu Ala Ala Tyr Tyr Glu Lys Lys Glu | 1037 | | | | | | | | |
| 152 | • | | | | | | | | | |
| | GCT TTT GCA AAG ACA ACT AAA CTA ACG CTC AAT GCA AAA AGT ATT CGC | 1085 | | | | | | | | |
| | Ala Phe Ala Lys Thr Thr Lys Leu Thr Leu Asn Ala Lys Ser Ile Arg | 1005 | | | | | | | | |
| 156 | | | | | | | | | | |
| | TGC TGT TTA CAT TAGAAATCAC TTCCAGCTTA CATCTTACAC GGTGTCTTAC | 1137 | | | | | | | | |
| | Cys Cys Leu His | | | | | | | | | |
| | 310 | | | | | | | | | |
| | AAATTCTGCT AGTCTGTAAG CTCCTTAAGA GTAGGGTTGT GCCTTATTCA ACTGCATACA | 1197 | | | | | | | | |
| 164 | TAGCTCCTAG CACAGTGCCT TATTCGGTAG GCATCTAAGC AAATTTCTTA AATTAATTAA | 1257 | | | | | | | | |
| 166 | TATATCTTTA AAGATATCAT ATTTTATGTA TGTAGCTTAT TCAAAGAAGT GTTTCCTATT | 1317 | | | | | | | | |
| 168 | TCTATATAGT TTATTATACA TGATACTTGG GTAGCTCAAC ATTCTTAATA AACAGCCTTT | 1377 | | | | | | | | |
| 170 GTATTCAGAA TATAAAATTG AAATAGATAT ATATAAAGTT AAAAAAAA | | | | | | | | | | |
| 173 | (2) INFORMATION FOR SEQ ID NO: 2: | | | | | | | | | |
| 175 | • • | | | | | | | | | |
| | 176 (A) LENGTH: 313 amino acids | | | | | | | | | |
| | 177 (B) TYPE: amino acid | | | | | | | | | |
| 178 | · · | | | | | | | | | |
| 181 (ix) FEATURE: | | | | | | | | | | |
| | 182 (A) NAME/KEY: Modified-site | | | | | | | | | |
| 183 | 183 (B) LOCATION: 83 | | | | | | | | | |
| | • • • | | | | | | | | | |
| 186 (ix) FEATURE: 187 (A) NAME/KEY: Modified-site | | | | | | | | | | |
| 188 | · · · | | | | | | | | | |
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| 192 | , <i>,</i> | | | | | | | | | |
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| 198 | (ix) FEATURE: | | | | | | | | | |
| 199 | 199 (A) NAME/KEY: Modified-site | | | | | | | | | |
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| 209 | (A) NAME/KEY: Modified-site | | | | | | | | | |

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| | | | | | | 17 | | | | | | | | | | |
|------------|---|--------|------------|-------|-------|----------|-------|--------|--------|-------|--------------|-------|-------|-------|-------|------|
| 210 | | | LOC | | | | ON: | /not | - n | Doto | ntia | 1 NI- | alve | nevi | atio | 'n |
| 211 | | (ט) |) OTE | iek i | NF OF | CIMA I I | .ON: | /1100 | .e- | FULE | :11 (1 (| 11 | gryc | .0571 | .ucic | ••• |
| | 212 site" | | | | | | | | | | | | | | | |
| 214 | | | | | | | | | | | | | | | | |
| 215 | 40 04 | | | | | | | | | | | | | | | |
| 216 | | | | | | | | | | | | | | | | |
| 217 | | | | | | | | | | | | | | | | |
| | 18 predicted to be involved in catalysis" | | | | | | | | | | | | | | | |
| 220 | | | | | | | | | | | | | | | | |
| 221 | 0.00 | | | | | | | | | | | | | | | |
| 222 | (B) LOCATION: 275278 | | | | | | | | | | | | | | | |
| 223 | | | | | | | | | | | | | | | | |
| 224 | 24 predicted to be involved in catalysis" | | | | | | | | | | | | | | | |
| 226 | 26 (ix) FEATURE: | | | | | | | | | | | | | | | |
| 227 | | | | | | | ≀e-si | .te | | | | | | | | |
| 228 | | (B |) LO | CATIO | N: 2 | 283. | . 289 | | | | | | | | | |
| 229 | | | | | | | ON: | | | 'Cons | sensı | ıs se | equer | nce | | |
| 230 | predicte | d to | be i | invo] | Lved | in o | catal | ysis | 3 " | | | | | | | |
| 233 | (xi) | SEQ | JENCI | E DES | SCRI | OITS | N: SI | EQ II | ON C | 2: | | | | | | |
| 235 | Met | Thr | Ser | Cys | His | Ile | Ala | Glu | Glu | His | Ile | Gln | Lys | Val | Ala | Ile |
| 236 | 1 | | | | 5 | | | | | 10 | | | | | 15 | |
| 238 | Phe | Gly | Gly | Thr | His | Gly | Asn | Glu | Leu | Thr | Gly | Val | Phe | Leu | Val | Lys |
| 239 | | - | • | 20 | | | | | 25 | | | | | 30 | | |
| 241 | His | Trp | Leu | Glu | Asn | Gly | Ala | Glu | Ile | Gln | Arg | Thr | Gly | Leu | Glu | Val |
| 242 | | • | 35 | | | _ | | 40 | | | | | 45 | | | |
| 244 | Lvs | Pro | Phe | Ile | Thr | Asn | Pro | Arg | Ala | Val | Lys | Lys | Cys | Thr | Arg | Tyr |
| 245 | -1- | 50 | | | | | 55 | _ | | | | 60 | | | | |
| 247 | Ile | Asp | Cvs | Asp | Leu | Asn | Arg | Ile | Phe | Asp | Leu | Glu | Asn | Leu | Gly | Lys |
| 248 | 65 | | -1- | | | 70 | _ | | | - | 75 | | | | | 80 |
| 250 | | Met | Ser | Glu | Asp | | Pro | Tyr | Glu | Val | Arg | Arg | Ala | Gln | Glu | Ile |
| 251 | 2,0 | | | | 85 | | | • | | 90 | _ | | | | 95 | |
| 253 | λςn | His | T.eu | Phe | Glv | Pro | Lys | Asp | Ser | Glu | Asp | Ser | Tyr | Asp | Ile | Ile |
| 254 | 11011 | | Lou | 100 | 1 | | -1- | | 105 | | - | | _ | 110 | | |
| 256 | Dhe | Asn | Len | | Asn | Thr | Thr | Ser | Asn | Met | Gly | Cys | Thr | Leu | Ile | Leu |
| 257 | The | . mpp | 115 | | | | | 120 | | | • | - | 125 | | | |
| 259 | Glu | Asn | Ser | Arσ | Asn | Asn | Phe | Leu | Ile | Gln | Met | Phe | His | Tyr | Ile | Lys |
| .260 | OIG | 130 | 001 | ***** | | | 135 | | | | | 140 | | - | | |
| 262 | Thr | | T.011 | Δla | Pro | Len | Pro | Cvs | Tvr | Val | Tvr | Leu | Ile | Glu | His | Pro |
| | | | пси | AIU | 110 | 150 | 110 | 0,10 | -1- | | 155 | | | | | 160 |
| 263 265 | 145 | LAII | Lvc | ጥህዮ | | | Thr | | | | | | | | | |
| 266 | 261 | пец | цуз | 1 7 1 | 165 | * 111 | | *** 9 | 002 | 170 | | -1- | - 1 | | 175 | • |
| | т1 с | C111 | Val | C137 | | Gln | Pro | G1n | Glv | | Leu | Ara | Ala | Asp | | Leu |
| 268 | 116 | GIU | Val | 180 | FIO | GIII | 110 | 0111 | 185 | | | 5 | | 190 | | |
| 269 | 3.00 | C1 n | Wot | | Two | Mot | Ile | T.37 C | | Δla | Len | Asp | Phe | | His | His |
| 271 | ASP | GTII | мес 195 | ита | пуз | me c | 116 | 200 | 11.1.0 | 111u | | | 205 | | | |
| 272 | Dh- | . 7.~- | | C1** | T *** | c1 | Phe | | Dro | Ctre | Δ 1 = | Tle | | Va 1 | Tvr | Lvs |
| 274 | hue | | GIU | ату | пур | GIU | 215 | 110 | 110 | Cys | 2.14 | 220 | | | -1- | _1 _ |
| 275 | + 3 | 210 | α1 | T | 17-7 | 7 c~ | Tyr | Dro | λνα | Δen | Glu | | Glv | Glu | Tle | Ala |
| 277 | | | GIU | гÀЗ | val | _ | тАт | FIO | лту | voh | 235 | ron | OTY | Jiu | -10 | 240 |
| 278 | 225 | 1 | | | | 230 | | | | | ∠ .) .) | | | | | 240 |

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DATE: 11/19/2001 TIME: 14:24:06

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Ala Ile Ile His Pro Asn Leu Gln Asp Gln Asp Trp Lys Pro Leu His
280
                                              250
281
                          245
         Pro Gly Asp Pro Met Phe Leu Thr Leu Asp Gly Lys Thr Ile Pro Leu
283
                                          265
284
         Gly Gly Asp Cys Thr Val Tyr Pro Val Phe Val Asn Glu Ala Ala Tyr
286
                                      280
287
                 275
         Tyr Glu Lys Lys Glu Ala Phe Ala Lys Thr Thr Lys Leu Thr Leu Asn
289
290
                                  295
                                                       300
         Ala Lys Ser Ile Arg Cys Cys Leu His
292
293
                              310
295 (2) INFORMATION FOR SEQ ID NO: 3:
297
         (i) SEQUENCE CHARACTERISTICS:
              (A) LENGTH: 313 amino acids
298
299
              (B) TYPE: amino acid
300
              (D) TOPOLOGY: linear.
        (ix) FEATURE:
303
304
              (A) NAME/KEY: Region
305
              (B) LOCATION: 6
              (D) OTHER INFORMATION: /note= "This is isoleucine in
306
307 human, valine in bovine. This is a very
308 conservative substitution."
310
        (ix) FEATURE:
311
              (A) NAME/KEY: Region
312
              (B) LOCATION: 9
              (D) OTHER INFORMATION: /note= "This is glutamic acid in
313
314 human, aspartic acid in bovine. This is a very
315 conservative substitution."
317
        (ix) FEATURE:
              (A) NAME/KEY: Region
318
319
              (B) LOCATION: 10
              (D) OTHER INFORMATION: /note= "This is histidine in human,
321 proline in bovine. This is a conservative
322 substitution."
        (ix) FEATURE:
326
              (A) NAME/KEY: Region
327
              (B) LOCATION: 12
328
              (D) OTHER INFORMATION: /note= "This is glutamine in human,
329
330 lysine in bovine. This is a very conservative
331 substitution."
333
        (ix) FEATURE:
334
              (A) NAME/KEY: Region
335
              (B) LOCATION: 38
              (D) OTHER INFORMATION: /note= "This is glycine in human,
336
337 serine in bovine. This is a very conservative
338 substitution."
340
        (ix) FEATURE:
341
              (A) NAME/KEY: Region
342
              (B) LOCATION: 39
              (D) OTHER INFORMATION: /note= "This is alanine in human,
343
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VERIFICATION SUMMARY

PATENT APPLICATION: US/09/965,807

DATE: 11/19/2001 TIME: 14:24:07

Input Set : N:\Crf3\RULE60\09965807.txt
Output Set: N:\CRF3\11192001\1965807.raw

L:33 M:220 C: Keyword misspelled or invalid format, [(A) APPLICATION NUMBER:] L:34 M:220 C: Keyword misspelled or invalid format, [(B) FILING DATE:] L:527 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 L:533 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 L:539 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 $L\!:\!542$ M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 $L\!:\!551$ $M\!:\!341$ W: (46) "n" or "Xaa" used, for SEQ ID#:3 L:563 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 L:569 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 L:572 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 L:576 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 L:579 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 L:585 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 L:845 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:12 L:884 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:14 L:887 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:14 L:911 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:15 L:932 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:16 L:1162 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:25 L:1190 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:26 L:1212 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:27